## **PCT**

## INTERNATIONAL PRELIMINARY EXAMINATION REPORT

(PCT Article 36 and Rule 70)

REC'D 3 1 OCT 2003

	Applicant's or agent's file reference REP06595WO			FOR FURTHER	ACTION	See Notification	n of Transmittal of International amination Report (Form PCT/IF	PEA/416)
1	matior T/GB		olication No. 375	International filing date 28.03.2003	(day/mont	h/year)	Priority date (day/month/year) 02.04.2002	)
	mation 7K14		ent Classification (IPC) or b	oth national classification	and IPC			
	licant K TH	ERAI	PEUTICS LTD. et al.					
1.	. This international preliminary examination report has been prepared by this International Preliminary Examining Authority and is transmitted to the applicant according to Article 36.							
2.	This REPORT consists of a total of 4 sheets, including this cover sheet.							
	This report is also accompanied by ANNEXES, i.e. sheets of the description, claims and/or drawings which have been amended and are the basis for this report and/or sheets containing rectifications made before this Authority (see Rule 70.16 and Section 607 of the Administrative Instructions under the PCT).							
	These annexes consist of a total of sheets.					·		
3.	This	repo	rt contains Indications rel	ating to the following i	tems:			
	1	$\boxtimes$	Basis of the opinion					
	!!		Priority					
	111				novelty, in	ventive step ar	nd industrial applicability	
	V V		Lack of unity of invention Reasoned statement un citations and explanation	nder Rule 66.2(a)(ii) w	ith regard	to novelty, inv	entive step or industrial app	olicability;
	VI		Certain documents cite		atement	,		
	VII		Certain defects in the ir		า	,		
	VIII		Certain observations or	n the international app	lication			
Date	of sub	missio	n of the demand		Date of c	ompletion of this	s report	<u> </u>
16.0	16.09.2003			30.10.2003				
Name	Name and mailing address of the international preliminary examining authority:				Authorize	ed Officer		150E AUX
European Patent Office D-80298 Munich Tel. +49 89 2399 - 0 Tx: 523656 epmu d Fax: +49 89 2399 - 4465				6 epmu d	Grossk Telephon	opf, R e No. +49 89 23	999-8714	<b>1</b> 100 100 100 100 100 100 100 100 100 1

# INTERNATIONAL PRELIMINARY EXAMINATION REPORT

International application No.

PCT/GB03/01375

I.	<b>Basis</b>	of the	report
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**Description, Pages** 

1. With regard to the **elements** of the international application (Replacement sheets which have been furnished to the receiving Office in response to an invitation under Article 14 are referred to in this report as "originally filed" and are not annexed to this report since they do not contain amendments (Rules 70.16 and 70.17)):

	1-	11	as originally filed			
	CI	aims, Numbers				
	1-6	6	as originally filed			
	Dr	awings, Sheets				
	1/4	-4/4	as originally filed			
S	equ	ence listing part of t	the description, pages:			
1,	as	originally filed				
2. With regard to the <b>language</b> , all the elements marked above were available or furnished to this Allanguage in which the international application was filed, unless otherwise indicated under this item.						
	The	ese elements were a	vailable or furnished to this Authority in the following language: , which is:			
		the language of a tr	ranslation furnished for the purposes of the international search (under Rule 23.1(b)).			
		the language of pub	plication of the international application (under Rule 48.3(b)).			
		the language of a tr Rule 55.2 and/or 55	anslation furnished for the purposes of international preliminary examination (under 3.3).			
3.	Wit inte	h regard to any <b>nucl</b> ernational preliminary	eotide and/or amino acid sequence disclosed in the international application, the examination was carried out on the basis of the sequence listing:			
	$\boxtimes$	contained in the inte	ernational application in written form.			
	$\boxtimes$	filed together with th	ne international application in computer readable form.			
	☐ furnished subsequently to this Authority in written form.					
	☐ furnished subsequently to this Authority in computer readable form.					
		The statement that to in the international a	the subsequently furnished written sequence listing does not go beyond the disclosure application as filed has been furnished.			
		The statement that t listing has been furn	the information recorded in computer readable form is identical to the written sequence ished.			
4.	The	amendments have r	esulted in the cancellation of:			
		the description,	pages:			
		the claims,	Nos.:			
		the drawings,	sheets:			

#### INTERNATIONAL PRELIMINARY **EXAMINATION REPORT**

International application No.

PCT/GB03/01375

5. 🗆	This report has been established as if (some of) the amendments had not been made, since they have been considered to go beyond the disclosure as filed (Rule 70.2(c)).
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(Any replacement sheet containing such amendments must be referred to under item 1 and annexed to this report.)

6. Additional observations, if necessary:

V. Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement

1-6

1. Statement

Novelty (N)

Yes: Claims No:

Claims

Inventive step (IS) Yes: Claims 1-6

> No: Claims

Industrial applicability (IA) Yes: Claims 1-6

> No: Claims

2. Citations and explanations

see separate sheet

### **EXAMINATION REPORT - SEPARATE SHEET**

#### Ad item V:

The quoted documents are:

(1) SOKER S ET AL: "Inhibition of Vascular Endothelial Growth Factor (VEGF)-induced endothelial cell proliferation by a peptide corresponding to the exon7-encoded domain of VEGF165" JOURNAL OF BIOLOGICAL CHEMISTRY. AMERICAN SOCIETY OF BIOLOGICAL CHEMISTS, BALTIMORE, MD, US, vol. 272, no. 50, 12 December 1997 (1997-12-12), pages 31582-31588

#### (2) WO 93/08473

It is known from D1 that peptides which are derived from exon-7 of VEGF are inhibitory for the VEGF binding to HUVECs (human umbilical vein endothelial cells) which express the NP-1 receptor.

However, according to D1 the terminal cysteine residue which is absent in the sequence of Claim 1 is considered to be essential for said inhibitory activity. Therefore, it must be regarded as surprising that peptides which no longer have this cysteine still have NP-1 antagonist activity.

Also D2 discloses a peptide which is nearly identical to the sequence of Claim 1 (see e.g. Claim 8 of D2). However, said peptide has an additional Y at the Nterminus. Moreover, it is neither cyclic nor is an inhibitory activity disclosed.

Therefore, novelty and inventive activity can be acknowledged.